

INTERNATIONAL SEARCH REPORT

International Application No
PCT/IB2004/002165

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7	C07K14/47	C12N15/12	A61K47/42	A61K38/17	A61P35/00
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According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, Sequence Search, WPI Data, BIOSIS, EMBASE

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 94/03597 A (DUCHESNE MARC ; TOCQUE BRUNO (FR); RHONE POULENC RORER SA (FR); SCHWEI) 17 February 1994 (1994-02-17) claims; figures 1-3; examples 2,3; sequence 5	1-5, 10-22, 29-31
X	& US 6 180 362 B1 (DUCHESNE MARC ET AL) 30 January 2001 (2001-01-30) claims; examples; sequence 6	1-5, 10-22, 29-31
X	WO 03/018630 A (FRENCH JULIET ; KENNEDY DEREK (AU); UNIV GRIFFITH (AU); HART DEREK (AU) 6 March 2003 (2003-03-06) page 50, line 23 - page 51, line 7; claims; figure 2; sequence 6	1-3, 10-12, 16-19, 21,29,31
		-/-

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

* Special categories of cited documents:

- *A* document defining the general state of the art which is not considered to be of particular relevance
- *E* earlier document but published on or after the International filing date
- *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- *O* document referring to an oral disclosure, use, exhibition or other means
- *P* document published prior to the international filing date but later than the priority date claimed

- *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- *Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- *&* document member of the same patent family

Date of the actual completion of the International search

21 October 2004

Date of mailing of the International search report

02/11/2004

Name and mailing address of the ISA

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Madruga, J

INTERNATIONAL SEARCH REPORT

International Application No

PCT/IB2004/002165

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DUCHESNE M ET AL: "Identification of the SH3 domain of GAP as an essential sequence for Ras_GAP-mediated signaling" SCIENCE, AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, US, vol. 259, 22 January 1993 (1993-01-22), pages 525-528, XP002092187 ISSN: 0036-8075 abstract; figure 2	1,2, 10-16, 29,31
X	YANG JIANG-YAN ET AL: "Antiapoptotic signaling generated by caspase-induced cleavage of RasGAP" MOLECULAR AND CELLULAR BIOLOGY, vol. 21, no. 16, August 2001 (2001-08), pages 5346-5358, XP002296743 ISSN: 0270-7306	1,2, 10-16, 26-31
Y	cited in the application page 5351, left-hand column, paragraph 2 - page 5352, right-hand column, paragraph 1; figures page 5354, left-hand column, paragraph 2 - right-hand column, paragraph 5	23-25
Y	LEBLANC VERONIQUE ET AL: "Ras-GTPase activating protein inhibition specifically induces apoptosis of tumour cells" ONCOGENE, vol. 18, no. 34, August 1999 (1999-08), pages 4884-4889, XP002296744 ISSN: 0950-9232 cited in the application the whole document	23-25
Y	SCHWARZE S ET AL: "In vivo protein transduction: delivery of a biologically active protein into the mouse" SCIENCE, AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, US, vol. 285, no. 5433, 3 September 1999 (1999-09-03), pages 1569-1572, XP002140133 ISSN: 0036-8075 cited in the application the whole document	6-9
A	YANG JIANG-YAN ET AL: "A subset of caspase substrates functions as the Jekyll and Hyde of apoptosis." EUROPEAN CYTOKINE NETWORK, vol. 13, no. 4, December 2002 (2002-12), pages 404-406, XP008035485 ISSN: 1148-5493 cited in the application the whole document	

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>YANG JIANG-YAN ET AL: "The RasGAP N-terminal fragment generated by caspase cleavage protects cells in a Ras/PI3K/Akt-dependent manner that does not rely on NFkappaB activation" JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 277, no. 17, 26 April 2002 (2002-04-26), pages 14641-14646, XP002296745 ISSN: 0021-9258 cited in the application the whole document</p> <p>-----</p>	
A	<p>FELDMANN P ET AL: "control of growth and differentiation by drosophila rasgap, a homolog of p120 ras-gtpase-activating protein" MOLECULAR AND CELLULAR BIOLOGY, AMERICAN SOCIETY FOR MICROBIOLOGY, WASHINGTON, US, vol. 19, no. 3, March 1999 (1999-03), pages 1928-1937, XP002255964 ISSN: 0270-7306 the whole document</p> <p>-----</p>	
A	<p>PRENDERGAST G C ET AL: "RAS REGULATORY INTERACTIONS: NOVEL TARGETS FOR ANTI-CANCER INTERVENTION" BIOESSAYS, CAMBRIDGE, GB, vol. 16, no. 3, March 1994 (1994-03), pages 187-191, XP001181926 ISSN: 0265-9247 the whole document</p> <p>-----</p>	
A	<p>WO 99/65947 A (KENIGSBERG MIREILLE ; DUCHESNE MARC (FR); BARLAT ISABELLE (FR); PARKER) 23 December 1999 (1999-12-23) cited in the application claims; examples</p> <p>-----</p>	

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Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. Claims Nos.: 21-28 because they relate to subject matter not required to be searched by this Authority, namely:
Rule 39.1(iv) PCT – Method for treatment of the human or animal body by therapy
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this International application, as follows:

1. As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- The additional search fees were accompanied by the applicant's protest.
 No protest accompanied the payment of additional search fees.

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International Application No
PCT/IB2004/002165

Patent document cited in search report		Publication date		Patent family member(s)		Publication date
WO 9403597	A	17-02-1994	FR CA EP WO JP US	2694296 A1 2141061 A1 0652952 A1 9403597 A1 7509609 T 6180362 B1		04-02-1994 17-02-1994 17-05-1995 17-02-1994 26-10-1995 30-01-2001
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WO 03018630	A	06-03-2003	WO EP	03018630 A1 1427754 A1		06-03-2003 16-06-2004
WO 9965947	A	23-12-1999	FR AU AU CA CN EP WO HU JP	2780062 A1 764139 B2 4151599 A 2331151 A1 1305497 T 1087997 A2 9965947 A2 0103152 A2 2002518001 T		24-12-1999 14-08-2003 05-01-2000 23-12-1999 25-07-2001 04-04-2001 23-12-1999 28-12-2001 25-06-2002

PCT REQUEST

Original (for SUBMISSION)

VIII-4-1	<p>Declaration: Inventorship (only for the purposes of the designation of the United States of America)</p> <p>Declaration of Inventorship (Rules 4.17(iv) and 51bis.1(a)(iv)) for the purposes of the designation of the United States of America.</p> <p>I hereby declare that I believe I am the original, first and sole (if only one inventor is listed below) or joint (if more than one inventor is listed below) inventor of the subject matter which is claimed and for which a patent is sought.</p> <p>This declaration is directed to the international application of which it forms a part (if filing declaration with application).</p> <p>I hereby declare that my residence, mailing address, and citizenship are as stated next to my name.</p> <p>I hereby state that I have reviewed and understand the contents of the above-identified international application, including the claims of said application. I have identified in the request of said application, in compliance with PCT Rule 4.10, any claim to foreign priority, and I have identified below, under the heading "Prior Applications", by application number, country or Member of the World Trade Organization, day, month, and year of filing, any application for a patent or inventor's certificate filed in a country other than the United States of America, including any PCT international application designating at least one country other than the United States of America, having a filing date before that of the application on which foreign priority is claimed.</p>
VIII-4-1- 1	Prior applications:

PCT REQUEST

Original (for SUBMISSION)

	<p>I hereby acknowledge the duty to disclose information that is known by me to be material to patentability as defined by 37 C.F.R. § 1.56, including for continuation-in-part applications, material information which became available between the filing date of the prior application and the PCT international filing date of the continuation-in-part application.</p> <p>I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.</p>
VIII-4-1- 1-1	Name (LAST, First)
VIII-4-1- 1-2	Residence: (city and either US State, if applicable, or country)
VIII-4-1- 1-3	Mailing address:
VIII-4-1- 1-4	Citizenship:
VIII-4-1- 1-5	Inventor's Signature: (if not contained in the request, or if declaration is corrected or added under Rule 26ter after the filing of the international application. The signature must be that of the inventor, not that of the agent)
VIII-4-1- 1-6	Date: (of signature which is not contained in the request, or of the declaration that is corrected or added under Rule 26ter after the filing of the international application)

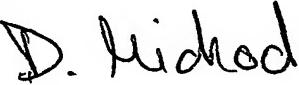
WIDMANN, Christian
 Lausanne, Switzerland
 Granette 5D CH-1010 Lausanne
 Switzerland
 CH



July 12, 2004

PCT REQUEST

Original (for SUBMISSION)

VIII-4-1- 2-1	Name (LAST, First)	YANG, Jiang-Yang
VIII-4-1- 2-2	Residence: (city and either US State, if applicable, or country)	Lausanne, Switzerland
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VIII-4-1- 2-4	Citizenship:	CN
VIII-4-1- 2-5	Inventor's Signature: (if not contained in the request, or if declaration is corrected or added under Rule 26ter after the filing of the international application. The signature must be that of the inventor, not that of the agent)	
VIII-4-1- 2-6	Date (of signature which is not contained in the request, or of the declaration that is corrected or added under Rule 26ter after the filing of the international application)	July 12, 2004
VIII-4-1- 3-1	Name (LAST, First)	MICHOD, David
VIII-4-1- 3-2	Residence: (city and either US State, if applicable, or country)	Lausanne, Switzerland
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VIII-4-1- 3-4	Citizenship:	CH
VIII-4-1- 3-5	Inventor's Signature: (if not contained in the request, or if declaration is corrected or added under Rule 26ter after the filing of the international application. The signature must be that of the inventor, not that of the agent)	
VIII-4-1- 3-6	Date (of signature which is not contained in the request, or of the declaration that is corrected or added under Rule 26ter after the filing of the international application)	July 12, 2004